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-Docket No. 17282 CIP

SEP 2 2 2003 F

## IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

PADA pplicant: Steward et al.	)	Group Art Unit: 1644
<u>-</u>	)	
Serial No.: 09/548,409	)	Examiner: Nolan, P.
Conf. No.: 7255	)	
<b>Filed:</b> April 13, 2000	)	
•	)	
For: Compositions and Methods	)	
For the Treatment of Pancreatitis	; )	
	1	

## **AMENDMENT C**

Commissioner for Patents Alexandria, VA 22313-1450

Dear Sir:

This communication is in reply to the Office Action mailed June 25, 2003. Applicants have the following comments.

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Docket: 17282 CIP Serial No. 09/548,409

STEWARD et al.

## **AMENDMENT**

- 1. (Previously amended) A composition for the treatment of acute pancreatitis in a mammal comprising, a first element comprising a binding element able to specifically bind a pancreatic acinar cell CCK receptor under physiological conditions, a second element comprising a translocation element derived from a clostridial neurotoxin heavy chain able to facilitate the transfer of a polypeptide across a vesicular membrane, and a third element comprising a therapeutic element derived from a clostridial neurotoxin light chain able, when present in the cytoplasm of a pancreatic cell, to inhibit enzymatic secretion by said pancreatic cell.
- 2. (Cancelled)
- 3. (Original) The composition of claim 1 wherein said therapeutic element will cleave a SNARE protein and cleavage of said SNARE protein inhibits said secretion.
- 4. (Original) The composition of claim 3 wherein said SNARE protein is selected from the group consisting of syntaxin, SNAP-25 and VAMP.
- 5. (Original) The composition of claim 2 wherein said therapeutic element will cleave a SNARE protein, wherein cleavage of said SNARE protein inhibits said secretion.
- 6. (Original) The composition of claim 5 wherein said SNARE protein is selected from the group consisting of syntaxin, SNAP-25 and VAMP.
- 7. (Original) The composition of claim 5 wherein said CCK receptor is the human CCK A receptor.
- 8. (Original) The composition of claim 7 wherein the binding element of said

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therapeutic polypeptide comprises a human CCK A amino acid sequence modified by the presence of a C-terminal amidated phenylalanine and a sulfated tyrosine at the position 7 residues from the carboxyl terminus.

- 9. (Original) The composition of claim 8 wherein said CCK A amino acid sequence comprises SEQ ID NO: 6.
- 10. (Original) The composition of claim 9 wherein said CCK A amino acid sequence comprises SEQ ID NO: 5.
- 11. (Original) The composition of claim 9 wherein said CCK A amino acid sequence comprises SEQ ID NO: 4.
- 12. (Original) The composition of claim 9 wherein said CCK A amino acid sequence comprises SEQ ID NO: 3.
- 13. (Original) The composition of claim 9 wherein said CCK A amino acid sequence comprises SEQ ID NO: 2.

14-19. (Cancelled)